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CERTIFICATE OF MAILING

I hereby certify that the foregoing document and all referenced attachments are being transmitted via Federal Express to Toni Hood, c/o RTIS, 3443 Carlin Springs Road, Falls Church, A 22041 this 20th day of December, 2004.

Registration No. 51,973

IN THE UNITED STATES PATENT AND TRADEMARK OFFICE

Application of: Wold, William S.M.

Group No.:

1632

Serial No.:

09/111,911

Atty. Docket No.: 66153-5587

Filed:

July 8, 1998

For:

INHIBITING APOPTOSIS

WITH ADENOVIRUS RID

PROTEIN

Examiner:

Ram R. Shukla, Ph.D.

Toni Hood c/o RTIS 3443 Carlin Springs Road Falls Church, VA 22041

PETITION UNDER 37 C.F.R. § 1.84(a)(2)

Applicant hereby petitions the United States Patent and Trademark Office to accept the enclosed color drawings.

REMARKS

In the present application, a color drawing of Figure 12 is necessary to clearly depict the location of Fas, LAMP-1, and their colocalization in rec700-infected A549 cells. The color green indicates Fas, red indicates LAMP-1, and yellow indicates colocalization of Fas and LAMP-1. If presented in black and white, the intensity of the dyes would not be clearly distinguishable from each other, such that the presence and location of Fas and LAMP-1 could not be clearly observed.

Figures 8, 18, 19 and 20 show flow cytometry tracings of A549 cells infected with various agents. The machine used to produce the tracings cannot print out the tracings with distinguishable black and white lines to show the resulting data. In order to present the same data in black and white, the figure would have to be redrawn using for example, dotted lines, dashed lines, or bold lines. However, the redrawing could not possibly capture the details of the machine-generated tracings, as depicted by the present Figures 8, 18, 19 and 20. Therefore, the color drawings are necessary.

Included in this petition is the (i) fee under 1.17(h); (ii) three (3) sets of each color drawing; and (iii) an amendment to the specification to insert the following language as the first paragraph of the brief description of the drawings:

The patent or application file contains at least one drawing executed in color. Copies of this patent or patent application publication with color drawing(s) will be provided by the Office upon request and payment of the necessary fee.

According to the MPEP 608.02, a black and white photocopy of the color drawings or color photographs has been waived. See 1246 O.G. 106 (May 22, 2001). Therefore, no black and white photocopy of the color drawings or color photographs are submitted herewith.

It is believed that the drawings are now in condition for reproduction and acceptance is respectfully requested.

Respectfully submitted,

Kimberly H. Lu, Reg. #51,973

Thompson Coburn LLP One US Bank Plaza

St. Louis, Missouri 63101

Telephone: 314-552-6000

Fax: 314-552-7000

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AMENDMENT UNDER 37 C.F.R. § 1.84

Responsive to a telephone conference with Toni Hood, please amend the above-identified application as follows:

Amendment to the Specification begins on page 2 of this paper.

Replacement Drawings are enclosed herewith per Examiner Hood's request.

Remarks begin on page 3 of this paper.

In the Specification:

Pursuant to 37 C.F.R. §121(b)(1), please amend the specification as follows:

(i) Please add the following <u>new</u> paragraph on page 6, line 4, after "<u>Brief Description of the Drawings</u>":

The patent or application file contains at least one drawing executed in color. Copies of this patent or patent application publication with color drawings will be provided by the Office upon request and payment of the necessary fee.

- (ii) Enclosed, please find the text of the above new paragraph with underlining to reflect the changes relative to the surrounding text;
- (iii) The full text of the added paragraph without underlining is shown above.

Remarks

Applicant is submitting herewith photographic drawings for Figs. 6, 9, 10, 11, 13, 14, 15, 21, 22, 23, 26, 29 and 30 to replace the original drawings. These drawings comply with 37 C.F.R. § 1.84(b)(1). Additionally, color replacement drawing sheets for Figs. 8, 12, 18, 19 and 20 are enclosed herewith, in triplicate. A petition to accept color drawings is also enclosed. It is believed that the drawings are now in condition for reproduction and acceptance is respectfully requested.

Respectfully submitted,

Kimberly H. Lu, Reg. #51,973

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dysfunction of apoptosis regulation; and the provision of compositions and methods for promoting tumor destruction in cancer patients.

Brief Description of the Drawings

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The patent or application file contains at least one drawing executed in color.

Copies of this patent or patent application publication with color drawing(s) will be provided by the Office upon request and payment of the necessary fee.

Figure 1 shows an alignment of the amino acid sequences of the death domains of the death receptor subfamily of TNFR proteins, with residues identical in more than 30% of sequences shaded black and residues conserved in more than 30% of sequences shaded in gray;

Figure 2 is a schematic representation of apoptosis mediated by death domain-containing members of the TNF receptor superfamily, with the death receptors Fas, TNFR1, TRAIL-R1, TRAIL-R2 and DR3 depicted by the bars on the extreme right and left sides of the figure, the ligands for these receptors indicated in parenthesis, and showing the association of the death receptors with intracellular proteins in the apoptotic singling cascade at the bottom of the figure;

Figure 3 is a schematic representation of a preferred RID complex showing one mature 14.5K polypeptide having an O-glycosylated residue in the extracellular (or lumenal) domain and an O-phosphorylated residue in the cytoplasmic domain, and two covalently-linked 10.4K polypeptides, one of which is an uncleaved, full-length form of 10.4K (10.4K-L) having two membrane-spanning regions (diagonal stripes) and the other a cleaved, short form of 10.4K (10.4K-S) with only one transmembrane region;

Figure 4 illustrates the amino acid sequences and various domains of preferred embodiments of the RID α and RID β polypeptides, showing in Fig. 4A-4B the long and short forms of the E3 10.4K polypeptides (RID α -L and RID α -S) from Ad serotype 2, Fig. 4C the pre-14.5K (RID β) polypeptide of Ad serotype 5, and in Fig. 4D the mature 14.5K (RID β) polypeptide of Ad serotype 5, with the signal sequences and transmembrane domains underlined and the asterisks indicating sites for disulfide linkage in RID α or for O-phosphorylation in RID β ;

Figure 5 is a schematic representation of a model for RID-induced internalization and degradation of Fas and TNFR1 death receptors, showing RID and the death receptor in the plasma membrane, entry of RID and the death receptor into endosomes, transport of these endosomes to lysosomes where the death receptor is degraded, and recycling of RID in endosomes to the cell surface, where it can internalize another death receptor molecule;

Docket Number: <u>66153-5587</u>

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In re Application of: Wold, William S.M.

Serial No.: 09/111,911 Filed: July 8, 1998

For: INHIBITING APOPTOSIS WITH ADENOVIRUS RID PROTEIN

Toni Hood c/o RTIS 3443 Carlin Springs Road Falls Church, VA 22041 Examiner: Ram R. Shukla, Ph.D.

Group Art Unit 1632

TRANSMITTAL LETTER

Enclosed herewith please find:

X Petition Under 37 C.F.R. § 1.84(a)(2).

X Amendment Under 37 C.F.R. § 1.84(a)(2).

X Photographic drawings for Figs. 6, 9, 10, 11, 13, 14, 15, 21, 22, 23, 26, 29 and 30.

X Replacement color drawing sheets (in triplicate) for Figs. 8, 12, 18, 19 and 20.

X A check in the amount of \$130.00 is attached. Please charge any deficiencies or credit any overpayment to Deposit Account 20-0823. A duplicate copy of this sheet is attached.

Respectfully submitted,

Kimberly H. Lu Reg. No. 51,973

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December 20, 2004

Publishing Division

DEC 2 7 200

Docket Number: <u>66153-5587</u>

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Toni Hood c/o RTIS 3443 Carlin Springs Road Falls Church, VA 22041 Examiner: Ram R. Shukla, Ph.D.

Group Art Unit 1632

Publishing Division

DEC 2 1, 2000

TRANSMITTAL LETTER

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Respectfully submitted,

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December 20, 2004